Please substitute the following set of claims for the pending claim set.

## **IN THE CLAIMS**

- 1-59. (canceled)
- 60. (Currently amended) A hypermutable, transgenic mouse wherein the germ and somatic cells of said mouse express comprise a transgenic polynucleotide encoding a dominant negative form of a PMS2 mismatch repair protein dominant negative allele of a PMS2 mismatch repair gene, wherein said dominant negative allele comprises a PMS2-134 allele.
- 61. (Currently amended) A hypermutable, transgenic mouse produced by a process comprising the steps of:

introducing a <u>transgenic</u> polynucleotide <u>comprising</u> encoding a <u>dominant negative form</u> of a <u>PMS2</u> mismatch repair protein a sequence encoding a <u>dominant negative allele of a PMS2</u> mismatch repair gene into a fertilized mouse egg, <u>wherein the dominant negative allele comprises a PMS2-134 allele</u>, whereby <u>said protein is expressed and said fertilized mouse egg</u> becomes hypermutable;

implanting the fertilized egg into a pseudopregnant female; and allowing said mouse egg to develop into a hypermutable, transgenic mouse.

62. (Currently amended) A method of making a hypermutable fertilized mouse egg comprising:

introducing into said fertilized mouse egg a <u>transgenic</u> polynucleotide <del>comprising</del> encoding a dominant negative form of a *PMS2* mismatch repair protein a sequence encoding a dominant negative allele of a *PMS2* mismatch repair gene, wherein the dominant negative allele comprises a *PMS2-134*-allele, whereby <u>said protein is expressed and</u> said fertilized mouse egg becomes hypermutable.

63-70. (canceled)

71. (Currently amended) A method for generating a mutation in a gene of interest comprising the steps of:

introducing a <u>transgenic</u> polynucleotide <u>comprising</u> <u>encoding</u> a dominant negative <u>allele</u> <u>form</u> of a *PMS2* mismatch repair <u>gene</u> <u>protein</u> into a fertilized mouse egg, <u>wherein the dominant</u> <u>negative allele comprises a *PMS2-134* allele, whereby <u>said protein is expressed and</u> the fertilized mouse egg becomes hypermutable;</u>

implanting the fertilized egg into a pseudopregnant female;

allowing said fertilized mouse egg to develop into a hypermutable, transgenic mouse; and testing the mouse to determine whether the gene of interest harbors a mutation.

- 72. (previously presented) The method of claim 71 wherein the step of testing comprises analyzing a nucleotide sequence of the gene of interest.
- 73. (previously presented) The method of claim 71 wherein the step of testing comprises analyzing mRNA transcribed from the gene of interest.
- 74. (previously presented) The method of claim 71 wherein the step of testing comprises analyzing a protein encoded by the gene of interest.
- 75. (previously presented) The method of claim 71 wherein the step of testing comprises analyzing the phenotype of the gene of interest.

76-80. (canceled)

81. (Currently amended) The method of claim 62 wherein the mismatch repair gene protein is human *PMS2* PMS2.

- 82. (Currently amended) The method of claim 81 wherein said <u>dominant negative form of a PMS2</u> mismatch repair <u>gene</u> <u>protein is encoded by a polynucleotide which</u> comprises a truncation mutation at codon 134 <del>as shown in</del> of SEQ ID NO:1.
- 83. (Currently amended) The method of claim 82 wherein the truncation mutation is a thymidine at nucleotide 424 of wild-type *PMS2* as shown in of SEQ ID NO:1.
- 84. (Currently amended) The hypermutable, transgenic mouse of claim 60 comprising a wherein the protein which consists of the first 133 amino acids of human PMS2.
- 85. (Currently amended) The hypermutable, transgenic mouse of claim 61 wherein the mismatch repair gene transgenic polynucleotide is human *PMS2*.
- 86. (Currently amended) The hypermutable, transgenic mouse of claim 61 wherein the transgenic polynucleotide dominant-negative allele comprises a truncation mutation at codon 134 of as shown in SEQ ID NO:1.
- 87. (Currently amended) The hypermutable, transgenic mouse of claim 86 wherein the truncation mutation is a thymidine at nucleotide 424 of wild-type *PMS2* of as shown in SEQ ID NO:1.
- 88. (Currently amended) The mouse of claim 60 wherein the mismatch repair gene protein is human PMS2 PMS2.
- 89. (Currently amended) The mouse of claim 88 wherein said <u>transgenic polynucleotide</u> mismatch repair-gene comprises a truncation mutation at codon 134 of as shown in SEQ ID NO:1.
- 90. (Currently amended) The mouse of claim 89 wherein the truncation mutation is a thymidine at nucleotide 424 of wild-type *PMS2* of as shown in SEQ ID NO:1.

- 91. (Currently amended) The method of claim 71 wherein the mismatch repair gene protein is human <u>PMS2 PMS2</u>.
- 92. (Currently amended) The method of claim 91 wherein said <u>transgenic polynucleotide</u> mismatch repair-gene comprises a truncation mutation at codon 134 of as shown in SEQ ID NO:1.
- 93. (Currently amended) The method of claim 92 wherein the truncation mutation is a thymidine at nucleotide 424 of wild-type *PMS2* of as shown in SEQ ID NO:1.
- 94. (Currently amended) The mouse of claim 61 comprising a wherein the protein which consists of the first 133 amino acid residues of human PMS2.
- 95. (Currently amended) The method of claim 62 wherein the <del>mouse egg comprises a</del> protein <del>which</del> consists of the first 133 amino acid residues of human PMS2.
- 96. (previously presented) The method of claim 71 wherein said mouse comprises a protein which consists of the first 133 amino acid residues of human PMS2.